```
Trying 3106016892...Open
Welcome to STN International! Enter x:x
LOGINID:ssspta1604dxj
PASSWORD:
TERMINAL (ENTER 1, 2, 3, OR ?):2
                     Welcome to STN International
                 Web Page URLs for STN Seminar Schedule - N. America
 NEWS 1
 NEWS 2 Dec 17 The CA Lexicon available in the CAPLUS and CA files
 NEWS 3 Feb 06 Engineering Information Encompass files have new names
 NEWS 4 Feb 16 TOXLINE no longer being updated
 NEWS 5 Apr 23 Search Derwent WPINDEX by chemical structure
 NEWS 6 Apr 23 PRE-1967 REFERENCES NOW SEARCHABLE IN CAPLUS AND CA
 NEWS 7 May 07 DGENE Reload
 NEWS EXPRESS May 23 CURRENT WINDOWS VERSION IS V6.0a,
              CURRENT MACINTOSH VERSION IS V5.0C (ENG) AND V5.0JB (JP),
              AND CURRENT DISCOVER FILE IS DATED 06 APRIL 2001
 NEWS HOURS
              STN Operating Hours Plus Help Desk Availability
 NEWS INTER
              General Internet Information
 NEWS LOGIN
              Welcome Banner and News Items
 NEWS PHONE
              Direct Dial and Telecommunication Network Access to STN
 NEWS WWW
             CAS World Wide Web Site (general information)
Enter NEWS followed by the item number or name to see news on that
specific topic.
 All use of STN is subject to the provisions of the STN Customer
  agreement. Please note that this agreement limits use to scientific
 research. Use for software development or design or implementation
  of commercial gateways or other similar uses is prohibited and may
  result in loss of user privileges and other penalties.
  * * * * * * * * * * * * * * * STN Columbus
                                          * * * * * * * * * * * * * * *
FILE 'HOME' ENTERED AT 12:24:52 ON 12 JUN 2001
=> file reg
COST IN U.S. DOLLARS
                                                 SINCE FILE
                                                                 TOTAL
                                                      ENTRY
                                                               SESSION
FULL ESTIMATED COST
                                                       0.15
                                                                 0.15
FILE 'REGISTRY' ENTERED AT 12:25:07 ON 12 JUN 2001
USE IS SUBJECT TO THE TERMS OF YOUR STN CUSTOMER AGREEMENT.
PLEASE SEE "HELP USAGETERMS" FOR DETAILS.
COPYRIGHT (C) 2001 American Chemical Society (ACS)
STRUCTURE FILE UPDATES:
                         10 JUN 2001 HIGHEST RN 340232-86-2
DICTIONARY FILE UPDATES: 10 JUN 2001 HIGHEST RN 340232-86-2
TSCA INFORMATION NOW CURRENT THROUGH January 11, 2001
  Please note that search-term pricing does apply when
  conducting SmartSELECT searches.
Structure search limits have been increased. See HELP SLIMIT
for details.
=> s amiodarone/cn
L1
            1 AMIODARONE/CN
=> d 11
```

ANSWER 1 OF 1 REGISTRY COPYRIGHT 2001 ACS

1951-25-3 REGISTRY

\$%^STN;HighlightOn= \*\*\*;HighlightOff=\*\*\* ;

T.1

RN

```
diodophenyl] - (9CI)
                          (CA INDEX NAME)
OTHER CA INDEX NAMES:
     Ketone, 2-butyl-3-benzofuranyl 4-[2-(diethylamino)ethoxy]-3,5-diiodophenyl
     (7CI, 8CI)
OTHER NAMES:
     2-Butyl-3-benzofuranyl p-[(2-diethylamino)ethoxy]-m,m-diiodophenyl ketone
CN
     2-Butyl-3-[3,5-diiodo-4-(2-diethylaminoethoxy)benzoyl]benzofuran
CN
CN
     2-n-Butyl-3',5'-diiodo-4'-N-diethylaminoethoxy-3-benzoylbenzofuran
CN
       ***Amiodarone***
CN
     Sedacoron
CN
     Sedacorone
FS
     3D CONCORD
MF
     C25 H29 I2 N O3
CI
     COM
LC
     STN Files:
                 AGRICOLA, AIDSLINE, ANABSTR, BEILSTEIN*, BIOBUSINESS, BIOSIS,
       BIOTECHNO, CA, CANCERLIT, CAOLD, CAPLUS, CASREACT, CBNB, CHEMCATS,
       CHEMLIST, CIN, CSCHEM, DDFU, DIOGENES, DRUGNL, DRUGPAT, DRUGU,
       DRUGUPDATES, EMBASE, HSDB*, IMSDIRECTORY, IPA, MEDLINE, MRCK*, NIOSHTIC,
       PHAR, PROMT, RTECS*, SPECINFO, TOXLINE, TOXLIT, USAN, USPATFULL, VETU
         (*File contains numerically searchable property data)
     Other Sources: EINECS**, WHO
         (**Enter CHEMLIST File for up-to-date regulatory information)
/ Structure 1 in file .gra /
            1225 REFERENCES IN FILE CA (1967 TO DATE)
              17 REFERENCES TO NON-SPECIFIC DERIVATIVES IN FILE CA
            1225 REFERENCES IN FILE CAPLUS (1967 TO DATE)
               5 REFERENCES IN FILE CAOLD (PRIOR TO 1967)
=> s dronedarone/cn
             1 DRONEDARONE/CN
L2
=> d 12
L2
     ANSWER 1 OF 1 REGISTRY COPYRIGHT 2001 ACS
RN
     141626-36-0 REGISTRY
CN
     Methanesulfonamide, N-[2-butyl-3-[4-[3-(dibutylamino)propoxy]benzoyl]-5-
     benzofuranyl] - (9CI) (CA INDEX NAME)
OTHER NAMES:
CN
      ***Dronedarone***
CN
     SR 33589
FS
     3D CONCORD
MF
     C31 H44 N2 O5 S
CI
     COM
SR
LC
                  ADISINSIGHT, BIOSIS, CA, CAPLUS, DDFU, DRUGNL, DRUGU,
       DRUGUPDATES, EMBASE, PROMT, TOXLIT, USAN, USPATFULL
     Other Sources:
                      WHO
/ Structure 2 in file .gra /
              20 REFERENCES IN FILE CA (1967 TO DATE)
               2 REFERENCES TO NON-SPECIFIC DERIVATIVES IN FILE CA
              20 REFERENCES IN FILE CAPLUS (1967 TO DATE)
=> file medline, embase, embal, ca, caplus, biosis'
COST IN U.S. DOLLARS
                                                  SINCE FILE
                                                                  TOTAL
                                                       ENTRY
                                                                SESSION
FULL ESTIMATED COST
                                                       11.22
                                                                  11.37
FILE 'MEDLINE' ENTERED AT 12:26:09 ON 12 JUN 2001
```

Methanone, (2-butyl-3-benzofuranyl)[4-[2-(diethylamino)ethoxy]-3,5-

CN

FILE 'EMBASE' ENTERED AT 12:26:09 ON 12 JUN 2001 COPYRIGHT (C) 2001 Elsevier Science B.V. All rights reserved. FILE 'EMBAL' ENTERED AT 12:26:09 ON 12 JUN 2001 COPYRIGHT (C) 2001 Elsevier Science B.V. All rights reserved. FILE 'CA' ENTERED AT 12:26:09 ON 12 JUN 2001 USE IS SUBJECT TO THE TERMS OF YOUR STN CUSTOMER AGREEMENT. PLEASE SEE "HELP USAGETERMS" FOR DETAILS. COPYRIGHT (C) 2001 AMERICAN CHEMICAL SOCIETY (ACS) FILE 'CAPLUS' ENTERED AT 12:26:09 ON 12 JUN 2001 USE IS SUBJECT TO THE TERMS OF YOUR STN CUSTOMER AGREEMENT. PLEASE SEE "HELP USAGETERMS" FOR DETAILS. COPYRIGHT (C) 2001 AMERICAN CHEMICAL SOCIETY (ACS) FILE 'BIOSIS' ENTERED AT 12:26:09 ON 12 JUN 2001 COPYRIGHT (C) 2001 BIOSIS(R) => s 11 or 12 'CN' IS NOT A VALID FIELD CODE 21282 L1 OR L2 => s cordarone or amiodarone or benzofuran or dronedarone 37729 CORDARONE OR AMIODARONE OR BENZOFURAN OR DRONEDARONE L4=> s 13 or 14 L5 37780 L3 OR L4 => s anionic surfactant L6 22946 ANIONIC SURFACTANT => s poloxamer? or polyethyoxylated castor oil? or ethoxylated polysorbate? or polyethylene hydr L7 45179 POLOXAMER? OR POLYETHYOXYLATED CASTOR OIL? OR ETHOXYLATED POLYSO RBATE? OR POLYETHYLENE HYDROXYSTEARATE? OR TWEEN? OR CREMOPHOR OR SOLUTROL => s 15 and 17 66 L5 AND L7 L8=> dup rem ENTER L# LIST OR (END):18 PROCESSING COMPLETED FOR L8 33 DUP REM L8 (33 DUPLICATES REMOVED) => s oral or tablet or capsule or gelatin or pill . L101486690 ORAL OR TABLET OR CAPSULE OR GELATIN OR PILL => s 19 and 110 T.11 4 L9 AND L10 => d 111 1-4 bib, ab, kwic L11 ANSWER 1 OF 4 MEDLINE 91346480 MEDLINE AN DN 91346480 PubMed ID: 2102806 TΙ Hemodynamic profile of \*\*\*amiodarone\*\*\* during acute and long-term administration in patients with ventricular dysfunction.

Cardiovascular Research Foundation Sticares, Rotterdam, The Netherlands.

ΑIJ

CS

Remme W J; van Hoogenhuyze D C

```
SO
     CARDIOSCIENCE, (1990 Sep) 1 (3) 169-76. Ref: 19
     Journal code: A5U; 9014943. ISSN: 1015-5007.
CY
DΤ
     Journal; Article; (JOURNAL ARTICLE)
     General Review; (REVIEW)
     (REVIEW, TUTORIAL)
LΑ
     English
FS
     Priority Journals
ΕM
     199110
ED
     Entered STN: 19911020
     Last Updated on STN: 19911020
     Entered Medline: 19911003
AΒ
     One of the potential adverse effects of anti-arrhythmic agents is an
     impairment of cardiac function as a result of their intrinsic negative
     inotropic properties. ***Amiodarone*** , in animals, also induces
     dose-related negative inotropic effects, in addition to coronary and
     systemic vasodilatation and slowing of the heart. Likewise, in most human
     studies, intravenous ***amiodarone***
                                             gives rise to early systemic and
     coronary vasodilatation, followed by a reduction in contractility.
     Depending on the relative impact of these opposing effects on the left
     ventricle, the changes in heart rate, cardiac output and left ventricular
     filling pressure are variable. Particularly in patients with pre-existing
     ventricular dysfunction, cardiac pump function is impaired further when
     relatively high dosages of ***amiodarone***
                                                   are used without its
              ***Tween*** 80. In contrast, fast bolus administrations, eg. 5
     solvent
            ***amiodarone*** in 5 minutes, result in an improvement of
     cardiac output, albeit at the expense of an increase in left ventricular
     filling pressure. The latter observation suggests that intravenous
       ***amiodarone***
                        should be given with caution in patients with heart
     failure and elevated left ventricular filling pressures. When given by
     mouth, ***amiodarone***
                                does not have significant hemodynamic effects,
     other than a moderate reduction in heart rate and, occasionally, in
     diastolic blood pressure. Cardiac pump function is not affected, even in
     patients with ventricular dysfunction or heart failure, in whom chronic
       ***oral***
                   administration of the drug is well tolerated.
TΤ
     Hemodynamic profile of
                             ***amiodarone***
                                                during acute and long-term
     administration in patients with ventricular dysfunction.
     . . . adverse effects of anti-arrhythmic agents is an impairment of
AΒ
     cardiac function as a result of their intrinsic negative inotropic
                  ***Amiodarone*** , in animals, also induces dose-related
     negative inotropic effects, in addition to coronary and systemic
     vasodilatation and slowing of the heart. Likewise, in most human studies,
                  ***amiodarone*** gives rise to early systemic and coronary
     vasodilatation, followed by a reduction in contractility. Depending on the
     relative impact of. . . are variable. Particularly in patients with
     pre-existing ventricular dysfunction, cardiac pump function is impaired
     further when relatively high dosages of
                                              ***amiodarone***
                          ***Tween*** 80. In contrast, fast bolus
     without its solvent
     administrations, eg. 5 mg/kg ***amiodarone*** in 5 minutes, result in
     an improvement of cardiac output, albeit at the expense of an increase in
     left ventricular filling pressure. The latter observation suggests that
                  ***amiodarone***
                                    should be given with caution in patients
     intravenous
     with heart failure and elevated left ventricular filling pressures. When
                      ***amiodarone***
                                         does not have significant hemodynamic
     given by mouth,
     effects, other than a moderate reduction in heart rate and, occasionally,
     in diastolic blood pressure. Cardiac pump function is not affected, even
     in patients with ventricular dysfunction or heart failure, in whom chronic
       ***oral***
                   administration of the drug is well tolerated.
CT
     Check Tags: Animal; Human
       *** Amiodarone: AD, administration & dosage***
       *** Amiodarone: AE, adverse effects***
       ****Amiodarone: TU, therapeutic use***
     *Arrhythmia: DT, drug therapy
     Dose-Response Relationship, Drug
     Heart Failure, Congestive: PP, physiopathology
     *Hemodynamics: DE, drug effects
     Myocardial.
RN
       ***1951-25-3 (Amiodarone) ***
L11
    ANSWER 2 OF 4 CA COPYRIGHT 2001 ACS
ΑN
     134:32972 CA
ΤI
     Porous drug matrixes containing polymers and sugars and methods of their
```

```
manufacture
ĮΝ
     Straub, Julie; Bernstein, Howard; Chickering, Donald E., III; Khatak,
     Sarwat; Randall, Greg
PΑ
     Acusphere, Inc., USA
SO
     PCT Int. Appl., 45 pp.
     CODEN: PIXXD2
DT
     Patent
LA
     English
FAN.CNT 1
                    KIND DATE
                                          APPLICATION NO.
                                                           DATE
     PATENT NO.
                           ------
                                           -----
     WO 2000072827 A2 20001207
WO 2000072827 A3 20010125
                                          WO 2000-US14578 20000525
                            20001207
PΙ
         W: AE, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CR, CU,
             CZ, DE, DK, DM, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL,
             IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA,
             MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI,
             SK, SL, TJ, TM, TR, TT, TZ, UA, UG, UZ, VN, YU, ZA, ZW, AM, AZ,
             BY, KG, KZ, MD, RU, TJ, TM
         RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY,
             DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ,
             CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG
PRAI US 1999-136323 P
                           19990527
     US 1999-158659 P
                            19991008
     US 1999-433486
                     Α
                            19991104
     US 2000-186310
                      Ρ
                            20000302
AB
     Drugs, esp. low aq. soly. drugs, are provided in a porous matrix form,
     preferably microparticles, which enhances dissoln. of the drug in aq.
     media. The drug matrixes preferably are made using a process that
     includes (i) dissolving a drug, preferably a drug having low aq. soly., in
     a volatile solvent to form a drug soln., (ii) combining at least one pore
     forming agent with the drug soln. to form an emulsion, suspension, or
     second solns., and (iii) removing the volatile solvent and pore forming
     agent from the emulsion, suspension, or second soln. to yield the porous
     matrix of drug. The pore forming agent can be either a volatile liq. that
     is immiscible with the drug solvent or a volatile solid compd., preferably
     a volatile salt. In a preferred embodiment, spray drying is used to
     remove the solvents and the pore forming agent. The resulting porous
     matrix has a faster rate of dissoln. following administration to a
     patient, as compared to non-porous matrix forms of the drug. In a
     preferred embodiment, microparticles of the porous drug matrix are
     reconstituted with an aq. medium and administered parenterally, or
     processed using std. techniques into tablets or capsules for ***oral***
     administration. Paclitaxel or docetaxel can be provided in a porous
     matrix form, which allows the drug to be formulated without solubilizing
     agents and administered as a bolus. For example, a nifedipine-loaded org.
     soln. was prepd. by dissolving 9.09 g of PEG 3350, 2.27 g of nifedipine,
     and 0.009 g of lecithin in 182 mL of methylene chloride. An aq. soln. was
     prepd. by dissolving 3.27 g of NH4HCO3 and 0.91 g of PEG 3350 in 1.82 mL
     of water. The aq. and org. solns. were homogenized and resulting emulsion
     was spray dried. A suspension of the porous nifedipine drug matrix was
     prepd. in 5% dextrose soln. at a concn. of 2.5 mg/mL. A bolus injection
     of the suspension was tolerated when administrated to dogs.
     Drugs, esp. low aq. soly. drugs, are provided in a porous matrix form,
AB
     preferably microparticles, which enhances dissoln. of the drug in aq.
     media. The drug matrixes preferably are made using a process that
     includes (i) dissolving a drug, preferably a drug having low aq. soly., in
     a volatile solvent to form a drug soln., (ii) combining at least one pore
     forming agent with the drug soln. to form an emulsion, suspension, or
     second solns., and (iii) removing the volatile solvent and pore forming
     agent from the emulsion, suspension, or second soln. to yield the porous
     matrix of drug. The pore forming agent can be either a volatile liq. that
     is immiscible with the drug solvent or a volatile solid compd., preferably
     a volatile salt. In a preferred embodiment, spray drying is used to
     remove the solvents and the pore forming agent. The resulting porous
     matrix has a faster rate of dissoln. following administration to a
     patient, as compared to non-porous matrix forms of the drug. In a
     preferred embodiment, microparticles of the porous drug matrix are
     reconstituted with an aq. medium and administered parenterally, or

tashsiques into tablets or capsules for ***oral***
     processed using std. techniques into tablets or capsules for
     administration. Paclitaxel or docetaxel can be provided in a porous
     matrix form, which allows the drug to be formulated without solubilizing
```

```
agents and administered as a bolus. For example, a nifedipine-loaded org.
soln. was prepd. by dissolving 9.09 g of PEG 3350, 2.27 g of nifedipine,
and 0.009 g of lecithin in 182 mL of methylene chloride. An aq. soln. was
prepd. by dissolving 3.27 g of NH4HCO3 and 0.91 g of PEG 3350 in 1.82 mL
of water. The aq. and org. solns. were homogenized and resulting emulsion
was spray dried. A suspension of the porous nifedipine drug matrix was
prepd. in 5% dextrose soln. at a concn. of 2.5 mg/mL. A bolus injection
of the suspension was tolerated when administrated to dogs.
drug solubilization polymer sugar porous matrix; microparticle
  ***oral***
             parenteral drug porous matrix
Drug delivery systems
     ***oral*** ; prepn. of porous matrixes contg. hydrophilic polymers
   and sugars for enhancement of drug dissoln.)
50-28-2, Estradiol, biological studies
                                        50-35-1, Thalidomide
                                                               50-99-7,
Dextrose, biological studies 52-53-9, Verapamil 53-03-2, Prednisone
                  57-63-6, Ethinyl estradiol 58-61-7, Adenosine,
55-98-1, Busulfan
                   59-92-7, Levodopa, biological studies
biological studies
67-97-0, Vitamin D3
                     67-97-0D, Vitamin D3, analogs
Medroxyprogesterone acetate 75-64-9, Erbumine, biological studies
77-36-1, Chlorthalidone
                        89-57-6, Mesalamine
                                              126-07-8, Griseofulvin
128-13-2, Ursodiol
                    298-46-4, Carbamazepine
                                              302-79-4, Tretinoin
321-64-2, Tacrine
                   363-24-6, Dinoprostone 437-38-7, Fentanyl
439-14-5, Diazepam 443-48-1, Metronidazole 518-28-5, Podofilox
                       846-49-1, Lorazepam
                                            ***1951-25-3***
745-65-3, Alprostadil
  ***Amiodarone***
                      3239-44-9, Dexfenfluramine 4759-48-2, Isotretinoin
                                        5593-20-4, Betamethasone
5534-09-8, Beclomethasone dipropionate
             9002-68-0, Follitropin
dipropionate
                                      9002-72-6, Growth hormone
9005-49-6, Enoxaparin, biological studies
                                          9007-12-9, Calcitonin
9041-93-4, Bleomycin sulfate
                             10238-21-8, Glyburide 11096-26-7,
                12629-01-5, Somatropin
                                        12633-72-6, Amphotericin
Erythropoietin
                      15307-79-6, Diclofenac sodium 15307-86-5,
13311-84-7, Flutamide
                                                            20830-75-5,
            15687-27-1, Ibuprofen
                                    18559-94-9, Albuterol
Diclofenac
        21256-18-8, Oxaprozin
                                21829-25-4, Nifedipine
                                                          22204-53-1,
Digoxin
         27203-92-5, Tramadol
                                28860-95-9, Carbidopa
                                                        28981-97-7,
Naproxen
            29094-61-9, Glipizide
                                    30516-87-1, Zidovudine
                                                             32986-56-4,
Alprazolam
           33069-62-4, Paclitaxel 34911-55-2, Bupropion
                                                             36505-84-7,
Tobramycin
           40391-99-9 41340-25-4, Etodolac 41575-94-4, Carboplatin
Buspirone
42399-41-7, Diltiazem 42924-53-8, Nabumetone 51022-70-9, Albuterol
                                 51773-92-3, Mefloquine hydrochloride
sulfate
         51333-22-3, Budesonide
                        54527-84-3, Nicardipine hydrochloride
54143-55-4, Flecainide
                        54965-21-8, Albendazole
54910-89-3, Fluoxetine
                                                 54965-24-1, Tamoxifen
         55268-75-2, Cefuroxime
                                  56124-62-0, Valrubicin
                                                           56180-94-0,
citrate
         59729-33-8, Citalopram 60142-96-3, Gabapentin
                                                           60205-81-4,
Acarbose
Ipratropium 63659-18-7, Betaxolol
                                     65277-42-1, Ketoconazole
                       66376-36-1, Alendronate
                                                 66852-54-8, Halobetasol
66085-59-4, Nimodipine
           69655-05-6, Didanosine
                                     70476-82-3, Mitoxantrone
propionate
               72432-03-2, Miglitol
                                      72509-76-3, Felodipine
hydrochloride
                        72956-09-3, Carvedilol
                                                  73384-59-5, Ceftriaxone
72558-82-8, Ceftazidime
                       75330-75-5, Lovastatin
73590-58-6, Omeprazole
                                                 75695-93-1, Isradipine
                       76095-16-4, Enalapril maleate
75847-73-3, Enalapril
                                                       76547-98-3,
            76824-35-6, Famotidine 76963-41-2, Nizatidine
Lisinopril
                                                             77883-43-3,
                                                          78628-80-5,
                   78246-49-8, Paroxetine hydrochloride
Doxazosin mesylate
Terbinafine hydrochloride 78755-81-4, Flumazenil
                                                   79517-01-4,
Octreotide acetate 79559-97-0, Sertraline hydrochloride
                                                           79794-75-5,
           79902-63-9, Simvastatin 80274-67-5, Metoprolol fumarate
Loratadine
81098-60-4, Cisapride
                      81103-11-9, Clarithromycin 82410-32-0,
              82752-99-6, Nefazodone hydrochloride 82834-16-0,
Ganciclovir
                                       83905-01-5, Azithromycin
              83799-24-0, Fexofenadine
Perindopril
                               84625-61-6, Itraconazole
83919-23-7, Mometasone furoate
                                                           85721-33-1,
                                        86541-74-4, Benazepril 87679-37-6, Trandolapril
               86386-73-4, Fluconazole
86541-75-5, Benazepril
Ciprofloxacin
hydrochloride
                                 91161-71-6, Terbinafine
89778-27-8, Toremifene citrate
                                                           91421-42-0,
                                    93957-54-1, Fluvastatin
           93413-69-5, Venlafaxine
Rubitecan
                         95233-18-4, Atovaquone
95058-81-4, Gemcitabine
                                                 97048-13-0,
                 97322-87-7, Troglitazone 98048-97-6, Fosinopril
Urofollitropin
                                       98319-26-7, Finasteride
98079-52-8, Lomefloxacin hydrochloride
99011-02-6, Imiquimod
                      99294-93-6, Zolpidem tartrate
                                                      100286-90-6,
Irinotecan hydrochloride 100986-85-4, Levofloxacin 103577-45-3,
              103628-48-4, Sumatriptan succinate 103775-10-6, Moexipril
Lansoprazole
104227-87-4, Famciclovir
                          104632-25-9, Pramipexole dihydrochloride 106463-17-6, Tamsulosin hydrochloride
106266-06-2, Risperidone
106685-40-9, Adapalene 107753-78-6, Zafirlukast 109889-09-0,
```

ST

ΙT

ΙT

```
110871-86-8, Sparfloxacin 111470-99-6, Amlodipine besylate
     Granisetron
     111974-72-2, Quetiapine fumarate 112809-51-5, Letrozole 113806-05-6,
     Olopatadine 114798-26-4, Losartan 114977-28-5, Docetaxel
     115956-12-2, Dolasetron 120014-06-4, Donepezil 124832-26-4,
     Valacyclovir 127779-20-8, Saquinavir 131918-61-1, Paricalcitol
     132539-06-1, Olanzapine 134308-13-7, Tolcapone 134678-17-4, Lamivudine
     137862-53-4, Valsartan 140678-14-4, Mangafodipir trisodium
     142373-60-2, Tirofiban hydrochloride 143011-72-7, Granulocyte
    colony-stimulating factor 144701-48-4, Telmisartan 145040-37-5,
    Candesartan cilexetil 147059-72-1, Trovafloxacin 147245-92-9,
    Glatiramer acetate 150378-17-9, Indinavir 154248-97-2, Imiglucerase
     154598-52-4, Efavirenz 155141-29-0, Rosiglitazone maleate 155213-67-5,
     Ritonavir 158966-92-8, Montelukast 159989-65-8, Nelfinavir mesylate
     161814-49-9, Amprenavir 162011-90-7, Rofecoxib 169590-42-5, Celecoxib
     171599-83-0, Sildenafil citrate
    RL: PEP (Physical, engineering or chemical process); THU (Therapeutic
    use); BIOL (Biological study); PROC (Process); USES (Uses)
        (prepn. of porous matrixes contg. hydrophilic polymers and sugars for
        enhancement of drug dissoln.)
     64-17-5, Ethanol, biological studies
                                          9003-43-4, Polyvinylpyrrolidine
     9005-65-6, ***Tween*** 80 25322-68-3, Polyethylene glycol
     26266-57-9, Span 40 106392-12-5, Pluronic F127
                                                     211733-74-3
     RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
        (prepn. of porous matrixes contg. hydrophilic polymers and sugars for
       enhancement of drug dissoln.)
L11
    ANSWER 3 OF 4 CA COPYRIGHT 2001 ACS
    133:301171 CA
    Compositions and methods for improved delivery of ionizable hydrophobic
     therapeutic agents
    Chen, Feng-jing; Patel, Manesh V.
    Lipocine, Inc., USA
    PCT Int. Appl., 99 pp.
    CODEN: PIXXD2
    Patent
    English
FAN.CNT 1
                                   APPLICATION NO. DATE
    PATENT NO. KIND DATE
    WO 2000059475 A1 20001012 WO 2000-US7342 20000316
        W: AE, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CR, CU,
            CZ, DE, DK, DM, DZ, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID,
            IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV,
            MA, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG,
            SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, UZ, VN, YU, ZA, ZW, AM,
            AZ, BY, KG, KZ, MD, RU, TJ, TM
        RW: GH, GM, KE, LS, MW, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY, DE,
            DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF,
            CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG
PRAI US 1999-287043
                     Α
                           19990406
     The present invention is directed to a pharmaceutical compn. including a
    hydrophobic therapeutic agent having at least one ionizable functional
     group, and a carrier. The carrier includes an ionizing agent capable of
     ionizing the functional group, a surfactant, and optionally solubilizers,
     triglycerides, and neutralizing agents. The invention further relates to
    a method of prepg. such compns. by providing a compn. of an ionizable
    hydrophobic therapeutic agent, an ionizing agent, and a surfactant, and
    neutralizing a portion of the ionizing agent with a neutralizing agent.
     The compns. of the invention are particularly suitable for use in
       ***oral*** dosage forms. A carrier contg. concd. phosphoric acid 0.025,
       ***Tween*** -20 0.3, Arlacel 186 0.2, sodium taurocholate 0.15, propylene
     glycol 0.3 g was formulated. Itraconazole was included in the carrier at
     30 mg/mL for testing the stability of the itraconazole soln. upon diln. in
     simulated gastric fluid.
RE.CNT 3
(1) Blair; US 4306981 A 1981 CA
(2) Hauer; US 5342625 A 1994 CA
(3) Story; US 4944949 A 1990 CA
    The present invention is directed to a pharmaceutical compn. including a
    hydrophobic therapeutic agent having at least one ionizable functional
    group, and a carrier. The carrier includes an ionizing agent capable of
```

IT

ΑN

ΤI

IN

PA

SO

DT

LA

ΡI

AB

RE

AB

ionizing the functional group, a surfactant, and optionally solubilizers, triglycerides, and neutralizing agents. The invention further relates to a method of prepg. such compns. by providing a compn. of an ionizable hydrophobic therapeutic agent, an ionizing agent, and a surfactant, and neutralizing a portion of the ionizing agent with a neutralizing agent. The compns. of the invention are particularly suitable for use in \*\*\*oral\*\*\* dosage forms. A carrier contg. concd. phosphoric acid 0.025, \*\*\*Tween\*\*\* -20 0.3, Arlacel 186 0.2, sodium taurocholate 0.15, propylene glycol 0.3 g was formulated. Itraconazole was included in the carrier at 30 mg/mL for testing the stability of the itraconazole soln. upon diln. in simulated gastric fluid. Castor oil RL: THU (Therapeutic use); BÍOL (Biological study); USES (Uses) (hydrogenated, ethoxylated, \*\*\*Cremophor\*\*\* RH 40; pharmaceutical compns. contg. hydrophobic therapeutic agents and carriers contg. ionizing agents and surfactants and triglycerides) Drug delivery systems \*\*\*oral\*\*\* ; pharmaceutical compns. contg. hydrophobic therapeutic agents and carriers contq. ionizing agents and surfactants and triglycerides) Drug delivery systems \*\*\*oral\*\*\* ; pharmaceutical compns. contg. hydrophobic (solns., therapeutic agents and carriers contq. ionizing agents and surfactants and triglycerides) 50-06-6, Phenobarbital, biological studies 50-21-5, biological studies 50-21-5D, Lactic acid, glycerides 50-44-2, Mercaptopurine 50-52-2, Thioridazine 50-53-3, Chlorpromazine, Amitriptyline biological studies 50-55-5, Reserpine 50-78-2 50-81-7, Ascorbic acid, biological studies 51-48-9, Levothyroxine, biological studies 51-52-5, Propylthiouracil 51-55-8, Atropine, biological studies 51-64-9, Dexamphetamine 52-86-8, Haloperidol 53-86-1, Indomethacin 54-05-7, Chloroquine 54-11-5, Nicotine 54-31-9 56-54-2, Quinidine 57-10-3, Palmitic acid, biological studies 57-11-4, Stearic acid, 57-27-2, Morphine, biological biological studies 57-22-7, Vincristine 57-41-0, Phenytoin 57-43-2, Amylobarbital 57-44-3, Barbital studies 57-47-6, Physostigmine 57-66-9, Probenecid 57-88-5, Cholesterol, biological studies 58-14-0, Pyrimethamine 58-25-3, Chlordiazepoxide 58-32-2, Dipyridamole 58-38-8, Prochlorperazine 58-39-9, Perphenazine 58-54-8, Ethacrynic acid 58-73-1, Diphenhydramine 58-94-6, 59-05-2, Methotrexate 59-66-5, Acetazolamide Chlorothiazide 59-96-1, Phenoxybenzamine 61-56-3, Sulthiame 61-68-7, Nitrofurazone Mefenamic acid 61-72-3, Cloxacillin 64-18-6, Formic acid, biological studies 64-19-7, Acetic acid, biological studies 64-77-7, Tolbutamide 65-85-0, Benzoic acid, biological studies 66-76-2, Dicumarol 66-79-5, Oxacillin 67-20-9, Nitrofurantoin 68-04-2, Sodium Citrate 68-11-1, Thioglycolic acid, biological studies 68-35-9, Sulfadiazine 69 - 23 - 8, 69-72-7, biological studies 69-93-2, Uric acid, Fluphenazine 72-69-5, Nortriptyline 72-44-6, Methaqualone biological studies 75-75-2, Methanesulfonic acid 76-57-3, Codeine 74-55-5, Ethambutol 76-74-4, Pentobarbital 76-99-3, Methadone 77-28-1, Butobarbital 77-36-1, Chlorthalidone 77-86-1, Tromethamine 77-92-9, biological studies 79-09-4, Propanoic acid, biological studies 79-10-7, Acrylic acid, biological studies 82-92-8, Cyclizine 83-68-1, Vitamin K6 83-70-5, Vitamin K5 83-89-6, Mepacrine 86-21-5, 83-69-2, Vitamin K7 86-35-1, Ethotoin 86-42-0, 86-22-6, Brompheniramine Pheniramine 87-69-4 89-57-6, Mesalamine 89-65-6, Isoascorbic acid Amodiaguine 90-82-4, Pseudoephedrine 90-84-6, Diethylpropion 94-20-2, 97-23-4, Dichlorophen 99-66-1, Valproic acid Chlorpropamide 101-31-5, Hyoscyamine 102-71-6, biological studies 104-15-4, p-Toluenesulfonic acid, biological studies 107-15-3, 1,2-Ethanediamine, biological studies 107-92-6, Butyric acid, biological studies 110-15-6, Butanedioic acid, biological studies 110-16-7, 2-Butenedioic acid (2Z)-, biological studies 110-17-8, Fumaric acid, biological studies 110-27-0, Isopropyl myristate 111-03-5, Glyceryl monooleate 111-62-6, Ethyl Oleate 111-90-0, Transcutol 112-80-1, Oleic acid, biological studies 113-15-5, Ergotamine 113-45-1, Methylphenidate 113-59-7, Chlorprothixene 113-92-8 114-07-8, Erythromycin 115-38-8,

121-44-8, biological

125-28-0, Dihydrocodeine

Methylphenobarbital 117-89-5, Trifluoperazine

124-04-9, Hexanedioic acid, biological studies

studies 122-09-8, Phentermine 122-20-3, Triisopropanolamine

125-53-1, Oxyphencyclimine 125-84-8, Aminoglutethimide 127-09-3, Sodium Acetate 127-33-3, Demeclocycline 127-69-5, Sulfafurazole

ΙT

ΙT

ΙT

IT

```
127-71-9, Sulfabenzamide
                         127-79-7, Sulfamerazine
                                                   128-13-2,
Ursodeoxycholic acid 128-37-0, Butylated Hydroxytoluene, biological
        129-03-3, Cyproheptadine 129-20-4, Oxyphenbutazone 130-95-0,
Quinine
          132-17-2, Benztropine 138-36-3, p-Bromophenylsulfonic acid
139-33-3, Edetate Disodium 141-43-5, biological studies 142-18-7,
Glyceryl monolaurate 142-91-6, Isopropyl palmitate 143-07-7, Lauric
acid, biological studies 144-11-6, Benzhexol 144-55-8, Sodium hydrogen
carbonate, biological studies 144-62-7, Ethanedioic acid, biological
         144-80-9, Sulfacetamide 144-83-2, Sulfapyridine 145-42-6,
Taurocholic acid, sodium salt 146-22-5, Nitrazepam 146-54-3,
               148-79-8, Thiabendazole 151-21-3, Sodium Dodecyl
Fluopromazine
Sulfate, biological studies 154-42-7, Thioguanine 190-39-6, Bisanthene
288-14-2, Isoxazole 298-57-7, Cinnarizine 299-42-3, Ephedrine
300-62-9, Amphetamine 302-79-4, Tretinoin 305-03-3, Chlorambucil
321-64-2, Tacrine 359-83-1, Pentazocine 361-37-5, Methysergide
364-62-5, Metoclopramide 389-08-2 396-01-0, Triamterene 404-86-4,
Capsaicin 437-38-7, Fentanyl 439-14-5, Diazepam 442-52-4, Clemizole
443-48-1, Metronidazole 446-86-6, Azathioprine 458-24-2, Fenfluramine
463-79-6, Carbonic acid, biological studies 471-34-1, Calcium carbonate,
biological studies 486-16-8, Carbinoxamine · 500-92-5, Proguanil 511-12-6, Dihydroergotamine 514-65-8, Biperiden 519-23-3, Ellipticine
522-00-9, Ethopropazine 523-87-5, Dimenhydrinate 525-66-6 526-95-4,
D-Gluconic acid 536-33-4, Ethionamide 537-21-3, Chlorproguanil
544-35-4, Ethyl linoleate 544-63-8, Myristic acid, biological studies
548-73-2, Droperidol 561-27-3, Diamorphine 564-25-0, Doxycycline
569-65-3, Meclozine 577-11-7, Docusate sodium 599-79-1, Sulfasalazine
603-50-9, Bisacodyl 604-75-1, Oxazepam 631-61-8, Ammonium Acetate
644-62-2, Meclofenamic acid 657-24-9, Metformin 668-94-0,
4,5-Diphenylimidazole 671-16-9, Procarbazine 723-46-6,
Sulfamethoxazole 738-70-5, Trimethoprim 739-71-9, Trimipramine
745-65-3, Alprostadil 768-94-5, Amantadine 846-49-1, Lorazepam
846-50-4, Temazepam 848-75-9, Lormetazepam 865-21-4, Vinblastine
911-45-5, Clomiphene 915-30-0, Diphenoxylate 961-71-7, Phenbenzamine
968-81-0, Acetohexamide 1134-47-0, Baclofen 1156-19-0, Tolazamide
1309-42-8, Magnesium hydroxide 1310-58-3, Potassium Hydroxide,
biological studies 1310-73-2, Sodium Hydroxide, biological studies
1327-43-1, Magnesium aluminum silicate 1330-80-9, Propylene glycol
        1333-28-4, Undecenoic acid 1335-30-4, Aluminum silicate
1336-21-6, Ammonium Hydroxide 1338-39-2, Sorbitan monolaurate
1338-41-6, Sorbitan monostearate 1338-43-8, Sorbitan monooleate
1400-61-9, Nystatin 1404-90-6, Vancomycin 1406-05-9, Penicillins
1508-75-4, Tropicamide 1553-60-2, Ibufenac 1622-61-3, Clonazepam
1622-62-4, Flunitrazepam 1812-30-2, Bromazepam ***1951-25-3***
  ***Amiodarone*** 1972-08-3, Dronabinol 2022-85-7, Flucytosine
2030-63-9, Clofazimine 2062-78-4, Pimozide
                                              2078-54-8, Propofol
2447-57-6, Sulfadoxine 2487-39-0, Vitamin K-S (II) 2515-61-9,
1,5-Diphenylpyrazoline 2609-46-3, Amiloride
                                               2709-56-0, Flupentixol
2898-12-6, Medazepam 2998-57-4, Estramustine
RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
   (pharmaceutical compns. contg. hydrophobic therapeutic agents and
   carriers contq. ionizing agents and surfactants and triglycerides)
                                                3239-44-9,
3056-17-5, Stavudine
                      3116-76-5, Dicloxacillin
                                           4117-33-3, Lysine Ethyl Ester
Dexfenfluramine
                  3737-09-5, Disopyramide
                                                  5002-47-1, Fluphenazine
4342-03-4, Dacarbazine
                        4759-48-2, Isotretinoin
           5036-02-2, Tetramisole 5051-62-7, Guanabenz 5104-49-4,
decanoate
               5306-85-4, Dimethyl Isosorbide 5588-33-0, Mesoridazine
Flurbiprofen
5633-20-5, Oxybutynin 5786-21-0, Clozapine
                                              6452-71-7, Oxprenolol
6493-05-6, Pentoxifylline
                           6506-37-2, Nimorazole
                                                  7087-68-5,
Diisopropylethylamine
                       7261-97-4, Dantrolene 7416-34-4, Molindone
7647-01-0, Hydrochloric Acid, biological studies
                                                 7664-38-2, Phosphoric
acid, biological studies 7664-38-2D, Phosphoric acid, esters
7664-93-9, Sulfuric acid, biological studies
                                             7681-93-8, Natamycin
7689-03-4, Camptothecin 7697-37-2, Nitric acid, biological studies
7778-53-2, Potassium Phosphate 8007-43-0, Sorbitan sesquioleate
8045-34-9, Pentaerythritol stearate 9002-92-0, Polyoxyethylene lauryl
        9002-93-1 9002-96-4, D-.alpha.-Tocopheryl polyethylene glycol
            9004-74-4, Methoxy polyethylene glycol 9004-95-9,
succinate
Polyethylene glycol cetyl ether 9004-98-2, Polyoxyethylene oleyl ether
9004-99-3, Myrj 51 9005-00-9, Polyoxyethylene stearyl ether 9005-08-7,
Polyethylene glycol distearate 9005-32-7, Alginic acid 9005-64-5, Polysorbate 20 9005-65-6, Polysorbate 80 9005-66-7, ***Tween***
           ***Tween*** 60 9007-48-1, Polyglyceryl oleate 9011-21-6
```

ΙT

```
9016-45-9
            9014-67-9, Aloxiprin
                                              9062-73-1, Polyethylene
glycol sorbitan laurate 9062-90-2, Polyethylene glycol sorbitan oleate
10034-85-2, Hydriodic acid 10035-10-6, Hydrobromic acid, biological
         10043-35-3, Boric acid, biological studies 10238-21-8
10262-69-8, Maprotiline
                         10457-90-6, Bromperidol 10540-29-1, Tamoxifen
11140-04-8, Imwitor 988
                         12633-72-6, Amphotericin 12772-47-3,
                         13292-46-1, Rifampin 13392-28-4, Rimantadine
Pentaerythritol oleate
           itol oleate 13292-46-1, Rifampin 13392-28-4, Rimantadine 13655-52-2, Alprenolol 14028-44-5, Amoxapine 14611-51-9,
13523-86-9
Selegiline 14808-79-8, Sulfate, biological studies 15307-86-5, Diclofenac 15574-96-6, Pizotifen 15676-16-1, Sulpiride 15686
                                                             15686-51-8,
Clemastine
            15686-71-2, Cephalexin
                                     15686-83-6, Pyrantel
                                                             15687-27-1,
           16110-51-3, Cromoglicic acid 16773-42-5, Ornidazole
Ibuprofen
17560-51-9, Metolazone 17617-23-1, Flurazepam 18016-80-3, Lysuride
18507-89-6, Decoquinate 18559-94-9, Albuterol 19216-56-9, Prazosin 19387-91-8, Tinidazole 19794-93-5, Trazodone 20594-83-6, Nalbuphine
                                                 20594-83-6, Nalbuphine
21187-98-4, Gliclazide 21256-18-8, Oxaprozin
                                                 21645-51-2, Aluminum
hydroxide, biological studies 21738-42-1, Oxamniquine
                                                          21829-25-4,
Nifedipine 22071-15-4, Ketoprofen 22131-79-9, Alclofenac
                                                               22204-53-1
22232-71-9, Mazindol 22494-42-4, Diflunisal 22882-95-7, Isopropyl
           22916-47-8, Miconazole 22994-85-0, Benznidazole
linoleate
23031-25-6, Terbutaline 23110-15-8, Fumagillin 23288-49-5, Probucol
23593-75-1, Clotrimazole 24219-97-4, Mianserin
                                                   25339-99-5, Sucrose
monolaurate 25523-97-1, Dexchlorpheniramine. 25614-03-3, Bromocriptine
25637-84-7, Glyceryl dioleate 25637-97-2, Sucrose dipalmitate
25812-30-0, Gemfibrozil
                         25953-19-9, Cefazolin 26097-80-3, Cambendazole
26171-23-3, Tolmetin 26266-57-9, Sorbitan monopalmitate
                                                            26266-58-0,
Sorbitan trioleate 26402-22-2, Glyceryl monocaprate
                                                       26402-26-6,
Glyceryl monocaprylate 26446-38-8, Sucrose monopalmitate
                                                             26658-19-5,
Sorbitan tristearate 26839-75-8, Timolol
                                            26912-41-4D, Polyethylene
                           27195-16-0, Sucrose distearate
glycol caprate, glycerides
                                                              27203-92-5,
           27220-47-9, Econazole 27321-96-6, Polyethylene glycol
Tramadol
              27638-00-2, Glyceryl dilaurate 28395-03-1, Bumetanide
cholesterol
                                              28981-97-7, Alprazolam
28657-80-9, Cinoxacin 28911-01-5, Triazolam
29094-61-9, Glipizide
                       29122-68-7, Atenolol 29679-58-1, Fenoprofen
29767-20-2, Teniposide 30299-08-2, Clinofibrate 30909-51-4,
Flupentixol decanoate 31431-39-7, Mebendazole 31692-85-0, Glycofurol
33419-42-0, Etoposide
                        33671-46-4, Clotiazepam 33940-98-6
Nikkol Decaglyn 1L 34580-13-7, Ketotifen 34911-55-2, Bupropion
36322-90-4, Piroxicam
                       36330-85-5, Fenbufen 36354-80-0, Glyceryl
             36531-26-7, Oxantel
dicaprylate
                                   36894-69-6, Labetalol
              37220-82-9, ARLACEL 186
                                      37318-31-3, Crodesta F-160
37321-62-3, Lauroglycol FCC
                             37517-30-9, Acebutolol
                                                       38194-50-2,
          38304-91-5, Minoxidil 38821-53-3, Cephradine
                                                            39366-43-3,
                              41340-25-4, Etodolac
Magnesium aluminum hydroxide
                                                      41859-67-0,
Bezafibrate
             42200-33-9, Nadolol 42399-41-7, Diltiazem
                                                            42766-91-6,
            43200-80-2, Zopiclone 43210-67-9, Fenbendazole
Nikkol DHC
50679-08-8, Terfenadine 51192-09-7, Nikkol TMGO 5
                                                      51264-14-3,
           51322-75-9, Tizanidine 51384-51-1, Metoprolol
                                                              51481-61-9,
Cimetidine
           51803-78-2 51938-44-4, Sorbitan sesquistearate
52081-33-1, Mitomycins 52468-60-7, Flunarizine
                                                   52504-24-2, Softigen
767
      52581-71-2, Volpo 3 52942-31-1, Etoperidone
                                                      53168-42-6, Myvacet
       53179-11-6, Loperamide
                                53230-10-7, Mefloquine
                                                         53716-50-0,
Oxfendazole
              53988-07-1, Glyceryl dicaprate
                                              54029-12-8, Ricobendazole
54143-55-4, Flecainide
                        54340-58-8, Meptazinol
                                                  54392-26-6, Sorbitan
monoisostearate
                  54910-89-3, Fluoxetine
                                           55142-85-3, Ticlopidine
                          55985-32-5, Nicardipine
                                                     57107-95-6
55268-74-1, Praziquantel
57307-93-4, Pentaerythritol caprylate 57801-81-7, Brotizolam
57808-66-9, Domperidone
                          58581-89-8, Azelastine 59467-70-8, Midazolam
                         60142-96-3, Gabapentin
                                                  60607-34-3, Oxatomide
59729-33-8, Citalopram
60719-84-8, Amrinone
                      61318-90-9, Sulconazole
                                                 61379-65-5, Rifapentine
61869-08-7
            62013-04-1, Dirithromycin
                                         62571-86-2, Captopril
63590-64-7, Terazosin
                      63675-72-9, Nisoldipine
                                                  64211-45-6, Oxiconazole
64221-86-9, Imipenem
                       64840-90-0, Eperisone
                                              64872-76-0, Butoconazole
65271-80-9, Mitoxantrone
                           65277-42-1, Ketocopazole
                                                      65899-73-2,
              66085-59-4, Nimodipine
                                       66357-35-5, Ranitidine
Tioconazole
67227-56-9, Fenoldopam 67352-02-7
                                     67915-31-5, Terconazole
68506-86-5, Vigabatrin 68844-77-9, Astemizole
                                                  68958-64-5, Polyethylene
                            68993-42-0D, Polyethylene glycol caprylate,
glycol glyceryl trioleate
glycerides
             69070-98-0
                        69756-53-2, Halofantrine
                                                    70458-96-7,
              71125-38-7, Meloxicam
                                     71486-22-1, Vinorelbine
Norfloxacin
72432-03-2, Miglitol
                     72509-76-3, Felodipine 72559-06-9, Rifabutin
72803-02-2, Darodipine 73590-58-6, Omeprazole 74011-58-8, Enoxacin
```

74103-06-3, Ketorolac 74191-85-8, Doxazosin RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses) (pharmaceutical compns. contg. hydrophobic therapeutic agents and carriers contg. ionizing agents and surfactants and triglycerides) ANSWER 4 OF 4 CA COPYRIGHT 2001 ACS 133:286507 CA Formulation arrays for screening Galakatos, Nicholas; Langer, Robert S.; Putnam, David A. Millennium Pharmaceuticals, Inc., USA PCT Int. Appl., 46 pp. CODEN: PIXXD2 Patent English FAN.CNT 2 PATENT NO. KIND DATE APPLICATION NO. DATE WO 2000059627 A1 20001012 WO 2000-US8589 20000331 W: AE, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CR, CU, CZ, DE, DK, DM, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, UZ, VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM RW: GH, GM, KE, LS, MW, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG PRAI US 1999-127755 19990405 P US 1999-146019 Ρ 19990728 Methods are described for high throughput combinatorial formulation in combination with nanotechnol. and microarrays to improve properties of materials used as components of, or in the manuf. or use of, health care products, consumer products, agricultural products, nutraceutical products, veterinary products, products for use in manufg. or processing industries, military applications, and research reagents. In particular, the bioavailability and pharmacokinetics of drugs, esp. small mol. pharmaceuticals, are optimized by making many new formulations and selecting those formulations based on phys. or chem. properties such as soly. in an aq. soln., without compromising selectivity or potency. Systems employing these technologies are described for rapid, systematic and cheap identification of optimal compns. for a desired purpose. New formulations can be prepd. and tested for bioequivalence to a formulation that is approved or com. available. Addnl., formulations can be initially optimized in vitro for their pharmacokinetics, such as absorption through \*\*\*oral\*\*\* prepn.), skin (for transdermal the gut (for an application), or mucosa (for nasal, buccal, vaginal or rectal formulation), soly., degrdn. or clearance by uptake into the reticuloendothelial system ("RES"), metab. or elimination, then tested in vivo. RE.CNT 9 (2) Gold, G; JOURNAL OF PHARMACEUTICAL SCIENCES 1964, V53(1), P52 CA (4) McFarland, E; TRENDS IN BIOTECHNOLOGY 1999, V17(3), P107 CA (5) Pokorny, V; WO 9840159 A 1998 CA (6) Rothbard, J; WO 9852614 A 1998 CA (8) Song, C; JOURNAL OF CONTROLLED RELEASE 1997, V45(2), P177 CA ALL CITATIONS AVAILABLE IN THE RE FORMAT Methods are described for high throughput combinatorial formulation in combination with nanotechnol. and microarrays to improve properties of materials used as components of, or in the manuf. or use of, health care products, consumer products, agricultural products, nutraceutical products, veterinary products, products for use in manufg. or processing industries, military applications, and research reagents. In particular, the bioavailability and pharmacokinetics of drugs, esp. small mol. pharmaceuticals, are optimized by making many new formulations and selecting those formulations based on phys. or chem. properties such as soly. in an aq. soln., without compromising selectivity or potency. Systems employing these technologies are described for rapid, systematic and cheap identification of optimal compns. for a desired purpose. New formulations can be prepd. and tested for bioequivalence to a formulation

that is approved or com. available. Addnl., formulations can be initially optimized in vitro for their pharmacokinetics, such as absorption through

L11

ΑN

ΤI

ΙN

PA

SO

DT

LA

PΙ

AΒ

RE

```
· application), or mucosa (for nasal, buccal, vaginal or rectal
     formulation), soly., degrdn. or clearance by uptake into the
     reticuloendothelial system ("RES"), metab. or elimination, then tested in
     vivo.
ΙT
     Drug delivery systems
         ( ***oral*** ; formulation arrays for screening)
     110-82-7D, Cyclohexane, ***benzofuran*** derivs. complexes 112-80-1,
     Oleic acid, biological studies 121-54-0, Benzethonium chloride
     151-21-3, Sodium dodecyl sulfate, biological studies 271-89-6D,
     ***Benzofuran*** , derivs., cyclohexane complexes 3287-99-8D,
Benzylammonium chloride, trialkyl derivs. 6106-24-7, Sodium tartrate
dihydrate 7585-39-9, .beta.-Cyclodextrin 8044-71-1, Cetrimide
     9000-01-5, Gum arabic 9002-89-5, Polyvinyl alcohol 9002-92-0, BRIJ 35
     9004-98-2, BRIJ 97 9004-99-3, Polyethylene glycol stearate 9005-65-6,
       ***TWEEN*** 80
                          25322-68-3, Polyethylene glycol 106392-12-5,
       ***POLOXAMER***
                           237
     RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
         (formulation arrays for screening)
=> d his
     (FILE 'HOME' ENTERED AT 12:24:52 ON 12 JUN 2001)
     FILE 'REGISTRY' ENTERED AT 12:25:07 ON 12 JUN 2001
Ll
              1 S AMIODARONE/CN
L2
               1 S DRONEDARONE/CN
     FILE 'MEDLINE, EMBASE, EMBAL, CA, CAPLUS, BIOSIS' ENTERED AT 12:26:09 ON
     12 JUN 2001
L3
           21282 S L1 OR L2
L4
           37729 S CORDARONE OR AMIODARONE OR BENZOFURAN OR DRONEDARONE
L5
           37780 S L3 OR L4
L6
          22946 S ANIONIC SURFACTANT
          45179 S POLOXAMER? OR POLYETHYOXYLATED CASTOR OIL? OR ETHOXYLATED POL
L7
\Gamma8
              66 S L5 AND L7
              33 DUP REM L8 (33 DUPLICATES REMOVED)
L9
        1486690 S ORAL OR TABLET OR CAPSULE OR GELATIN OR PILL
L10
L11
               4 S L9 AND L10
=>
---Logging off of STN---
=>
Executing the logoff script...
=> LOG Y
COST IN U.S. DOLLARS
                                                    SINCE FILE
                                                                     TOTAL
                                                          ENTRY
                                                                   SESSION
FULL ESTIMATED COST
                                                          87.32
                                                                      98.69
DISCOUNT AMOUNTS (FOR QUALIFYING ACCOUNTS)
                                                    SINCE FILE
                                                                     TOTAL
                                                          ENTRY
                                                                   SESSION
CA SUBSCRIBER PRICE
                                                          -1.68
                                                                     -1.68
```

STN INTERNATIONAL LOGOFF AT 12:42:10 ON 12 JUN 2001

the gut (for an \*\*\*oral\*\*\*, prepn.), skin (for transdermal